

EXAMPLE 63**5'-O-DMT-2'-deoxy-2'-methylene-5-methyl uridine-3'-(2-cyanoethyl-N,N-diisopropyl) phosphoramidite**

[0241] 2'-Deoxy-2'-methylene-3',5'-O-(tetraisopropyl disiloxane-1,3,diyl)-5-methyl uridine is synthesized following the procedures reported for the corresponding uridine derivative (Hansske, F.; Madej, D.; Robins, M.J. *Tetrahedron* (1984) **40**, 125; Matsuda, A.; Takenusi, K.; Tanaka, S.; Sasaki, T.; Ueda, T. *J. Med. Chem.* (1991) **34**, 812; See also Cory, A. H.; Samano, V.; Robins, M. J.; Cory, J. G. 2'-Deoxy-2'-methylene derivatives of adenosine, guanosine, tubercidin, cytidine and uridine as inhibitors of L1210 cell growth in culture. *Biochem. Pharmacol.* (1994), 47(2), 365-71.)

[0242] It is treated with IM TBAF in THF to give 2'-deoxy-2'-methylene-5-methyl uridine. It is dissolved in pyridine and treated with DMT-Cl and stirred to give the 5'-O-DMT-2'-deoxy-2'-methylene-5-methyl uridine. This compound is treated with 2-cyanoethyl-N,N-diisopropyl phosphoramidite and diisopropylaminotetrazolide. In a similar manner the corresponding N-6 benzoyl adenosine, N-4-benzoyl cytosine, N-2-isobutyryl guanosine phosphoramidite derivatives are synthesized.

EXAMPLE 63**Synthesis of 3'-O-4'-C-methylenribonucleoside**

[0243] 5'-O-DMT-3'-O-4'-C-methylene uridine and 5-methyl uridine are synthesized and phosphorylated according to the procedure of Obika *et al.* (Obika *et al. Bioorg. Med. Chem. Lett.* (1999) **9**, 515-158). The amidites are incorporated into oligonucleotides using the protocols described above.

EXAMPLE 64**Synthesis of 2'-methylene phosphoramidites**

[0244] 5'-O-DMT-2'-(methyl)-3'-O-(2-cyanoethyl-N,N-diisopropylamine)-5-methyluridine-phosphoramidite, 5'-O-DMT-2'-(methyl)-N-6-benzoyl adenosine (3'-O-2-cyanoethyl-N,N-diisopropylamino) phosphoramidite, 5'-O-DMT-2'-(methyl)-N2-isobutyryl guanosine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidite and 5'-O-DMT-2'-(methyl)-N-4-benzoyl cytidine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidites were obtained by the phosphitylation of the corresponding nucleosides. The nucleosides were synthesized according to the procedure described by Iribarren, Adolfo M.; Cicero, Daniel O.; Neuner, Philippe J. Resistance to degradation by nucleases of (2'S)-2'-deoxy-2'-C-methyloligonucleotides, novel potential antisense probes. *Antisense Res. Dev.*, (1994), 4(2), 95-8; Schmit, Chantal; Bevierre, Marc-Olivier; De Mesmaeker, Alain; Altmann, Karl-Heinz. "The effects of 2'- and 3'-alkyl substituents on oligonucleotide hybridization and stability". *Bioorg. Med. Chem. Lett.* (1994), 4(16), 1969-74.

[0245] The phosphitylation is carried out by using the bisamidite procedure.

EXAMPLE 65**Synthesis of 2'-S-methyl phosphoramidites**

[0246] 5'-O-DMT-2'-S-(methyl)-3'-O-(2-cyanoethyl-N,N-diisopropylamine)-5-methyl uridine-phosphoramidite, 5'-O-DMT-2'-S-(methyl)-N-6-benzoyl adenosine (3'-O-2-cyanoethyl-N,N-diisopropylamino) phosphoramidite, 5'-O-DMT-2'-S-(methyl)-N2-isobutyryl guanosine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidite and 5'-O-DMT-2'-S-(methyl)-N-4-benzoyl cytidine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidites were obtained by the phosphitylation of the corresponding nucleosides. The nucleosides were synthesized according to the procedure described by Fraser *et al.* (Fraser, A.; Wheeler, P.; Cook, P.D.; Sanghvi, Y.S. *J. Heterocycl. Chem.* (1993) 31, 1277-1287). The phosphitylation is carried out by using the bisamidite procedure.

EXAMPLE 66**Synthesis of 2'-O-methyl-β-D-arabinofuranosyl compounds**

[0247] 2'-O-Methyl-β-D-arabinofuranosyl-thymidine containing oligonucleotides were synthesized following the procedures of Gotfredson *et. al.* (Gotfredson, C.H. *et. al. Tetrahedron Lett.* (1994) **35**, 6941-6944; Gotfredson, C.H. *et. al. Bioorg. Med. Chem.* (1996) **4**, 1217-1225). 5'-O-DMT-2'-ara-(O-methyl)-3'-O-(2-cyanoethyl-N,N-diisopropylamine)-5-methyl uridine-phosphoramidite, 5'-O-DMT-2'-ara-(O-methyl)-N-6-benzoyl adenosine (3'-O-2-cyanoethyl-N,N-diisopropylamino) phosphoramidite, 5'-O-DMT-2'-ara-(O-methyl)-N2-isoburytyl guanosine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidite and 5'-O-DMT-2'-ara-(O-methyl)-N-4-benzoyl cytidine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidites are obtained by the phosphitylation of the corresponding nucleosides. The nucleosides are synthesized according to the procedure described by Gotfredson, C.H. *et. al. Tetrahedron Lett.* (1994) **35**, 6941-6944; Gotfredson, C.H. *et. al. Bioorg. Med. Chem.* (1996) **4**, 1217-1225. The phosphitylation is carried out by using the bisamidite procedure.

EXAMPLE 67**Synthesis of 2'-fluoro-β-D-arabinofuranosyl compounds**

[0248] 2'-Fluoro-β-D-arabinofuranosyl oligonucleotides are synthesized following the procedures by Kois, P. *et al.*, Nucleosides Nucleotides **12**, 1093, 1993 and Damha *et al.*, J. Am. Chem. Soc., **120**, 12976, 1998 and references cited therein. 5'-O-DMT-2'-ara-(fluoro)-3'-O-(2-cyanoethyl-N,N-diisopropylamine)-5-methyl uridine-phosphoramidite, 5'-O-DMT-2'-ara-(fluoro)-N-6-benzoyl adenosine (3'-O-2-cyanoethyl-N,N-diisopropylamino) phosphoramidite, 5'-O-DMT-2'-ara-(fluoro)-N2-isoburytyl guanosine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidite and 5'-O-DMT-2'-ara-(fluoro)-N-4-benzoyl cytidine-3'-O-(2-cyanoethyl-N,N-diisopropylamino) phosphoramidites are obtained by the phosphitylation of the corresponding nucleosides. The nucleosides are synthesized according to the procedure described by Kois, P. *et*